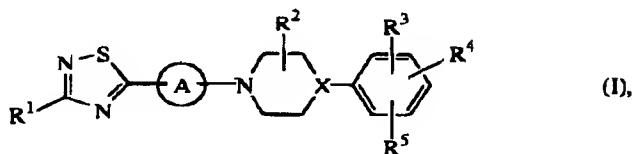


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JAB-1669

Amendments to the Claims:

## I. (Previously Amended) A compound of formula (I),



the *N*-oxide forms, the pharmaceutically acceptable acid addition salts and stereochemically isomeric forms thereof, wherein

X is N;

R<sup>1</sup> is hydrogen, C<sub>1</sub>-<sub>6</sub>alkyl, C<sub>1</sub>-<sub>6</sub>alkyloxy, C<sub>1</sub>-<sub>6</sub>alkylthio, amino, mono- or di(C<sub>1</sub>-<sub>6</sub>alkyl)amino, Ar<sup>1</sup>, Ar<sup>1</sup>NH-, C<sub>3</sub>-<sub>6</sub>cycloalkyl, hydroxymethyl or benzyloxymethyl;

R<sup>2</sup> is hydrogen, C<sub>1</sub>-<sub>6</sub>alkyl, amino, aminocarbonyl, mono- or di(C<sub>1</sub>-<sub>6</sub>alkyl)amino, C<sub>1</sub>-<sub>6</sub>alkyloxycarbonyl, C<sub>1</sub>-<sub>6</sub>alkylcarbonylamino, hydroxy or C<sub>1</sub>-<sub>6</sub>alkyloxy;

R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are each independently selected from hydrogen, halo, C<sub>1</sub>-<sub>6</sub>alkyl, C<sub>1</sub>-<sub>6</sub>alkyloxy, trifluoromethyl, nitro, amino, cyano, azido, C<sub>1</sub>-<sub>6</sub>alkyloxyC<sub>1</sub>-<sub>6</sub>alkyl, C<sub>1</sub>-<sub>6</sub>alkylthio, C<sub>1</sub>-<sub>6</sub>alkyloxycarbonyl or Het<sup>1</sup>;

is Ar<sup>2</sup> or Het<sup>2</sup>;

Ar<sup>1</sup> is phenyl; phenyl substituted with 1, 2 or 3 substituents each independently selected from halo, C<sub>1</sub>-<sub>6</sub>alkyl, C<sub>1</sub>-<sub>6</sub>alkyloxy, trihalomethyl, amino or nitro;



Ar<sup>2</sup> is

substituted with 1, 2 or 3

substituents each independently selected from halo, C<sub>1</sub>-<sub>6</sub>alkyl, C<sub>1</sub>-<sub>6</sub>alkyloxy, trihalomethyl, amino or nitro;

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Het<sup>1</sup> is a monocyclic heterocycle selected from oxazolyl, isoxazolyl, oxadiazolyl, thiazolyl, isothiazolyl, thiadiazolyl or oxazolinyl; and each monocyclic heterocycle may optionally be substituted on a carbon atom with C<sub>1-4</sub>alkyl; and

Het<sup>2</sup> is a monocyclic heterocycle selected from thiadiazolyl, pyridinyl, pyrimidinyl or pyrazinyl; and each monocyclic heterocycle may optionally be substituted on a carbon atom with 1 or 2 substituents each independently selected from halo, C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxy, nitro or trifluoromethyl.

2. (Previously Amended) A compound according to claim 1 wherein R<sup>1</sup> is hydrogen, C<sub>1-4</sub>alkyl, amino or di(C<sub>1-6</sub>alkyl)amino; R<sup>2</sup> is hydrogen; R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are each independently selected from hydrogen, halo, C<sub>1-6</sub>alkyl, C<sub>1-6</sub>alkyloxy, trifluoromethyl, nitro or C<sub>1-6</sub>alkyloxycarbonyl.

3. (Previously Amended) A compound according to claim 1 wherein R<sup>1</sup> is hydrogen, C<sub>1-4</sub>alkyl or di(C<sub>1-4</sub>alkyl)amino; R<sup>2</sup> is hydrogen; R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are each independently selected from hydrogen, halo, C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxy or trifluoromethyl; and the bivalent radical —— is Ar<sup>2</sup> or Het<sup>2</sup> wherein Ar<sup>2</sup> is phenyl and Het<sup>2</sup> is thiadiazolyl, pyridinyl, pyrimidinyl or pyrazinyl.

4. (Previously Amended) A compound according to claim 1 wherein R<sup>1</sup> is methyl, R<sup>2</sup> is hydrogen, R<sup>3</sup> and R<sup>4</sup> are hydrogen and R<sup>5</sup> is trifluoromethyl.

5. (Currently Amended) A compound according to claim 1 wherein the compound is 1-[4-(3-methyl-1,2,4-thiadiazol-5-yl)phenyl]-4-[3-(trifluoromethyl)phenyl]-piperazine; or

1-[5-(3-methyl-1,2,4-thiadiazol-5-yl)-2-pyridinyl]-4-[3-(trifluoromethyl)phenyl]-piperazine; a stereoisomeric form, or a pharmaceutically acceptable acid addition salt, or an N-oxide thereof.

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6. (Previously Amended) A composition comprising a pharmaceutically acceptable carrier, and as active ingredient a therapeutically effective amount of a compound as claimed in claim 1.

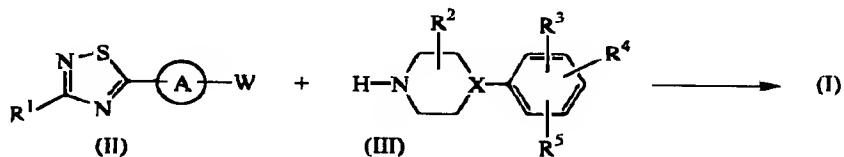
7. (Previously Cancelled).

8. (Previously Cancelled).

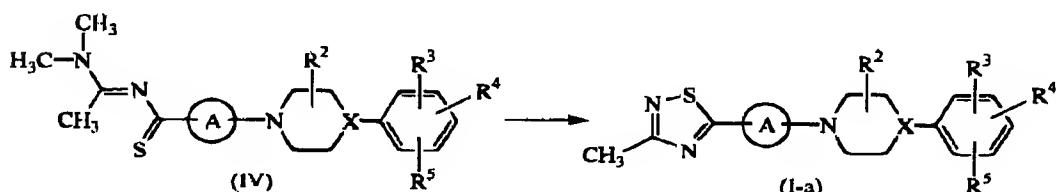
9. (Previously Cancelled).

10. (Currently Amended) A process of preparing a compound as claimed in claim 1, wherein

a) an intermediate of formula (II) is reacted with an intermediate of formula (III) in a reaction-inert solvent and, optionally in the presence of a suitable base;



b) an intermediate of formula (IV) is treated with hydroxylamino-O-sulfonic acid in a reaction-inert solvent, in the presence of a suitable base, thereby yielding compounds of formula (I-a), defined as compounds of formula (I) wherein  $\text{R}^1$  is methyl;



wherein in the above reaction schemes the radicals  $\text{X}$ ,  $\text{R}^1$ ,  $\text{R}^2$ ,  $\text{R}^3$ ,  $\text{R}^4$ ,  $\text{R}^5$  and

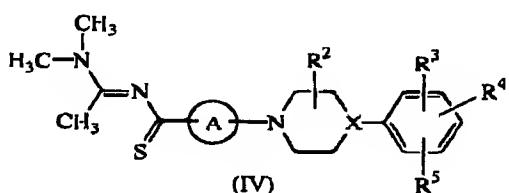
$\text{A}$  are as defined in claim 1, and  $\text{W}$  is an appropriate leaving group;

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c) or, a compound of formula (I) is converted into another compound of formula (I) by art-known group transformation reactions; or if desired; a compound of formula (I) is converted into a pharmaceutically acceptable acid addition salt, or conversely, an acid addition salt of a compound of formula (I) is converted into a free base form with alkali; and, if desired, preparing stereochemically isomeric forms thereof.

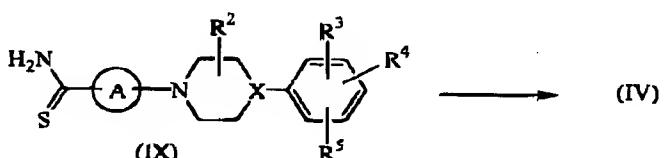
### 11. A compound of formula (IV),



an acid addition salt, a *N*-oxide form or a stereochemically isomeric form thereof, wherein  $X$ ,  $R^2$ ,  $R^3$ ,  $R^4$ ,  $R^5$  and the bivalent radical  $\text{—}\text{A}\text{—}$  are as defined in claim 1.

12. (Currently Amended) A process of preparing a compound of formula (IV) as claimed in claim 1011, wherein

a) an intermediate of formula (IX) is treated with *N,N*-dimethylacetamide dimethyl acetal in a reaction-inert solvent, thereby yielding a compound of formula (IV):



b) or, a compound of formula (IV) is converted into another compound of formula (IV) by art-known group transformation reactions; or if desired; a compound of formula (IV) is converted into an acid addition salt, or conversely, an acid addition salt of a compound of formula (IV) is converted into a free base form with alkali; and, if desired, preparing stereochemically isomeric forms thereof.

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13. A method of treating angiogenesis dependent disorders comprising administering to a host in need thereof an effective amount of a compound of claim 1.

14. A method of treating angiogenesis dependent disorders comprising administering to a host in need thereof an effective amount of a compound of claim 2.

15. A method of treating angiogenesis dependent disorders comprising administering to a host in need thereof an effective amount of a compound of claim 3.

16. A method of treating angiogenesis dependent disorders comprising administering to a host in need thereof an effective amount of a compound of claim 4.

17. A method of treating angiogenesis dependent disorders comprising administering to a host in need thereof an effective amount of a compound of claim 5.

18. (Previously Amended) A compound according to claim 2 wherein R<sup>1</sup> is hydrogen, C<sub>1-4</sub>alkyl or di(C<sub>1-4</sub>alkyl)amino; R<sup>2</sup> is hydrogen; R<sup>3</sup>, R<sup>4</sup> and R<sup>5</sup> are each independently selected from hydrogen, halo, C<sub>1-4</sub>alkyl, C<sub>1-4</sub>alkyloxy or trifluoromethyl; and the bivalent radical  is Ar<sup>2</sup> or Het<sup>2</sup> wherein Ar<sup>2</sup> is phenyl and Het<sup>2</sup> is thiadiazolyl, pyridinyl, pyrimidinyl or pyrazinyl.

19. (Previously Amended) A compound according to claim 2 wherein R<sup>1</sup> is methyl, R<sup>2</sup> is hydrogen, R<sup>3</sup> and R<sup>4</sup> are hydrogen and R<sup>5</sup> is trifluoromethyl.

20. (Previously Amended) A compound according to claim 3 wherein R<sup>1</sup> is methyl, R<sup>2</sup> is hydrogen, R<sup>3</sup> and R<sup>4</sup> are hydrogen and R<sup>5</sup> is trifluoromethyl.

21-37. (Cancelled).

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38. (Currently Amended) A method of treating angiogenesis dependent disorders comprising administering to a host in need thereof an effective amount of

1-[4-(3-methyl-1,2,4-thiadiazol-5-yl)phenyl]-4-[3-(trifluoromethyl)phenyl]-piperazine;

or

1-[5-(3-methyl-1,2,4-thiadiazol-5-yl)-2-pyridinyl]-4-[3-(trifluoromethyl)phenyl]-piperazine; a stereoisomeric form, or a pharmaceutically acceptable acid addition salt, or an N-oxide thereof.